

CLAIMS

- 1 1. A diagnostic method comprising the steps of:
 - 2 (a) examining an *in-vivo* tissue sample using an optical signal detection system adapted to
 - 3 automatically assign a classification to each of a plurality of regions of the tissue sample;
 - 4 (b) creating an overlay map visually indicating the classifications assigned to the regions of
 - 5 the tissue sample; and
 - 6 (c) displaying the overlay map to facilitate identification of suspect portions of the tissue
 - 7 sample.
- 1 2. The method of claim 1, wherein step (c) comprises superimposing the overlay map onto
- 2 an image of the tissue sample.
- 1 3. The method of claim 2, wherein the image of the tissue sample is a reference image.
- 1 4. The method of claim 3, wherein the tissue sample comprises cervical tissue and the
- 2 reference image is a colposcopic image.
- 1 5. The method of claim 2, wherein step (c) comprises displaying the overlay map
- 2 superimposed onto a real-time colposcopic image of the tissue sample.
- 1 6. The method of claim 1, wherein step (c) comprises projecting the overlay map onto the
- 2 tissue sample.
- 1 7. The method of claim 1, wherein steps (a) through (c) are performed during the course of
- 2 a single patient visit.
- 1 8. The method of claim 1, wherein the displaying step is performed substantially
- 2 contemporaneously with the examining step.

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1 9. The method of claim 1, wherein the classification in step (a) comprises a tissue-class
2 probability.

1 10. The method of claim 1, wherein the classifications assigned to the regions of the tissue
2 sample comprise at least one of the categories selected from the group consisting of necrotic,
3 CIN 2/3, NED, and indeterminate.

1 11. The method of claim 1, further comprising the steps of:

2 (d) identifying at least one suspect portion of the tissue sample; and

3 (e) marking the at least one suspect portion.

1 12. The method of claim 11, wherein step (e) comprises marking the at least one suspect
2 portion using at least one of an endogenous agent and an exogenous agent.

1 13. The method of claim 11, wherein step (e) comprises marking at least one suspect portion
2 using at least one of a photobleaching technique and a photoactivation technique.

1 14. The method of claim 11, wherein step (e) comprises marking the at least one suspect
2 region for follow-up examination.

1 15. The method of claim 11, wherein step (e) comprises marking the at least one suspect
2 region for treatment.

1 16. The method of claim 11, wherein steps (a) through (e) are performed during a single
2 patient visit.

1 17. The method of claim 1, further comprising the steps of:

2 (d) identifying at least one suspect portion of the tissue sample; and

3 (e) excising tissue from the at least one suspect portion for biopsy.

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1 18. The method of claim 1, further comprising the steps of:

2 (d) identifying at least one suspect portion of the tissue sample; and

3 (e) treating the at least one suspect portion.

1 19. The method of claim 18, wherein step (e) comprises performing at least one of:

2 photodynamic therapy, cryotherapy, and direct chemical treatment.

1 20. The method of claim 18, wherein step (e) comprises performing photodynamic therapy

2 using at least one photosensitive agent.

1 21. The method of claim 20, wherein the at least one photosensitive agent comprises at least

2 one of an exogenous agent and an endogenous agent.

1 22. The method of claim 20, wherein the at least one photosensitive agent comprises at least

2 one of a hematoporphyrin, a phthalocyanine, and a chlorin.

1 23. The method of claim 20, wherein the at least one photosensitive agent comprises at least

2 one of dihematoporphyrin, 5-aminolevulinic acid, protoporphyrin IX, temoporfin, and meso-

3 tetrahydroxyphenylchlorin.

1 24. The method of claim 18, wherein step (e) comprises removing tissue from the at least one

2 suspect portion using laser ablation.

1 25. The method of claim 18, wherein steps (a) through (e) are performed during a single

2 patient visit.

1 26. A system for detection of suspect portions of a tissue sample, the system comprising:

2 (a) an optical signal detection apparatus adapted to obtain at least one optical signal from

3 each of a plurality of regions of a tissue sample;

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- 4 (b) a memory that stores code defining a set of instructions;
- 5 (c) a processor that executes the instructions thereby to:
- 6 (i) identify a characteristic of each of the plurality of regions based at least in part on
- 7 the at least one optical signal; and
- 8 (ii) define an overlay map for visually indicating the characteristics identified in step
- 9 (i); and
- 10 (d) a display for facilitating identification of suspect portions of the tissue sample according
- 11 to the overlay map.

1 27. The system of claim 26, wherein the optical signal detection apparatus comprises

2 illuminating optics for illuminating each of the plurality of regions of the tissue sample and

3 collecting optics for receiving at least one optical signal from each of the plurality of regions.

1 28. The system of claim 27, wherein the illuminating optics comprise a white light source

2 and an ultraviolet light source and wherein the at least one optical signal comprises a

3 fluorescence spectrum and at least one reflectance spectrum.

1 29. The system of claim 28, wherein the at least one optical signal comprises a fluorescence

2 spectrum and two reflectance spectra.

1 30. The system of claim 27, wherein the optical signal detection apparatus comprises optics

2 for obtaining a sequence of images of the tissue sample useful for motion compensation.

1 31. The system of claim 27, wherein each of the plurality of regions has a dimension of about

2 a millimeter.

1 32. The system of claim 31, wherein each of the plurality of regions has a diameter of about

2 0.7 mm.

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1 33. The system of claim 27, wherein the optical signal detection apparatus is adapted to
2 complete a cervical scan in less than about 30 seconds.

1 34. The system of claim 33, wherein the optical signal detection apparatus is adapted to
2 complete a cervical scan in less than about 15 seconds.

1 35. The system of claim 26, wherein the display shows the overlay map superimposed onto
2 an image of the tissue sample.

1 36. The system of claim 35, wherein the image of the tissue sample is a reference image.

1 37. The system of claim 35, wherein the tissue sample comprises cervical tissue and the
2 reference image is a colposcopic image.

1 38. The system of claim 35, wherein the display shows the overlay map superimposed onto a
2 real-time colposcopic image of the tissue sample.

1 39. The system of claim 35, wherein the display can be viewed through a viewfinder.

1 40. The system of claim 35, wherein the display is a surface upon which the overlay map can
2 be projected.

1 41. The system of claim 26, wherein the display is a projection of the overlay map onto the
2 tissue sample.

1 42. The system of claim 26, further comprising a tool for marking a suspect region of the
2 tissue sample.

1 43. The system of claim 26, further comprising an instrument for excising tissue from a
2 suspect portion of the tissue sample for biopsy.

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1 44. The system of claim 26, further comprising an instrument for treating tissue from a
2 suspect portion of the tissue sample.

1 45. The system of claim 26, further comprising a laser for ablating tissue within a suspect
2 portion of the tissue sample.

1 46. The system of claim 26, further comprising a light source for performing photodynamic
2 therapy to treat the suspect portions of the tissue sample.

1 47. An optical signal detection apparatus for performing a cervical scan, the apparatus
2 comprising:

3 (a) illuminating optics for sequentially illuminating each of a plurality of regions of a tissue
4 sample, wherein the illuminating optics comprise a white light source and an ultraviolet light
5 source; and

6 (b) collecting optics for receiving at least one optical signal from each of the plurality of
7 regions, wherein the at least one optical signal comprises a fluorescence spectrum and at least
8 one reflectance spectrum, wherein each of the plurality of regions has a dimension of about a
9 millimeter, and wherein the optical signal detection apparatus is adapted to complete a cervical
10 scan in less than about 30 seconds.

1 48. The apparatus of claim 47, wherein the optical signal detection apparatus is adapted to
2 complete a cervical scan in less than about 15 seconds.

1 49. The apparatus of claim 47, wherein the plurality of regions of the tissue sample
2 comprises from about 100 to about 1000 regions of the tissue sample.

1 50. The apparatus of claim 49, wherein the plurality of regions of the tissue sample
2 comprises about 500 regions.

1 51. The apparatus of claim 47, comprising a light source for performing photodynamic
2 therapy.

1 52. A method for identifying a characteristic of each of a plurality of regions of a tissue
2 sample, the method comprising the steps of:

3 (a) obtaining at least one optical signal from each of a plurality of regions of a tissue sample
4 following application of a contrast agent and at least one of an optical probe and a biological-
5 responsive probe to the tissue sample; and

6 (b) automatically identifying a characteristic of each of the plurality of regions based at least
7 in part on the at least one optical signal.

1 53. The method of claim 52, wherein the optical probe comprises a spectroscopic enhancer.

1 54. The method of claim 52, wherein the biological-responsive probe comprises telomerase.

1 55. The method of claim 52, wherein step (b) comprises applying at least one of an optical
2 probe and a biological-responsive probe to the tissue sample in order to detect at least one
3 member selected from the group consisting of collagen, porphyrin, FAD, and NADH.

1 56. The method of claim 52, further comprising the steps of:

2 (e) creating an overlay map visually indicating the characteristics identified in step (d); and

3 (f) displaying the overlay map to facilitate identification of suspect portions of the tissue
4 sample.

1 57. The method of claim 52, wherein the at least one optical signal comprises a fluorescence
2 spectrum and at least one reflectance spectrum, and wherein step (b) comprises identifying a
3 characteristic of a subset of the plurality of regions based at least in part on a fluorescent
4 biomarker signal.